

LASIC et al.

Serial No.: 09/771,151

Filed: January 26, 2001

For: LIPOSOMES CONTAINING AN ENTRAPPED COMPOUND IN SUPERSATURATED SOLUTION

3. **(Amended)** The method of claim 1, wherein selecting the liposomes comprises [includes] selecting liposomes that have a liposome size of between about 60 nm to about 1000 nm.
4. **(Amended)** The method of claim 1, wherein selecting the liposomes comprises [includes] preparing liposomes having an entrapped compound at liposome size intervals between about 60 to about 1000 nm and analyzing the liposomes for the presence or absence of a precipitated compound.
5. **(Amended)** The method of claim 1, wherein selecting the liposomes comprises [includes] preparing liposomes having an entrapped compound at liposome size intervals between about 60 nm to about 1000 nm and analyzing the liposomes for the presence or absence of a precipitated compound.
6. **(Amended)** The method of claim 1, wherein the entrapping comprises [includes] preparing a solution of lipids.
7. **(Amended)** The method of claim 6, wherein the preparing comprises [includes] preparing a solution of lipids that comprises [include] a lipid derivatized with a hydrophilic polymer.
8. **(Amended)** The method of claim 6, wherein the preparing comprises [includes preparing] a solution of lipids effective to form a rigid lipid bilayer.
9. **(Amended)** The method of claim 1, further comprises [including] removing from an external liposome suspension medium the condition selected to maintain the drug above the room temperature solubility limit.

LASIC et al.

Serial No.: 09/771,151

Filed: January 26, 2001

For: LIPOSOMES CONTAINING AN ENTRAPPED COMPOUND IN SUPERSATURATED SOLUTION

11. A liposome composition comprising:

a suspension of liposomes composed of a vesicle-forming lipid, and

a compound entrapped in the liposomes, wherein the compound prior to entrapment is maintained in the liposomes in a supersaturated state.

12. The composition of claim 11, wherein the compound exhibits a two-fold increase in aqueous solubility in response to a condition selected from the group consisting of: (i) increasing solvent temperature, (ii) adding a co-solvent, and (iii) changing solvent pH.

13. The composition of claim 11, wherein the liposomes have a liposome size of between about 60 nm to about 1000 nm.

14. **(Amended)** The composition of claim 11 [1], wherein the liposomes further comprise a lipid derivatized with a hydrophilic polymer chain.

15. **(Amended)** The composition of claim 11 [1], wherein the liposomes comprise saturated vesicle-forming phospholipids.

16. A method for preparing liposomes comprising:

preparing an aqueous concentrated solution of a compound suitable for entrapment in an internal aqueous compartment of the liposomes;

hydrating a lipid film or lipid solution with a concentrated solution of the compound to form liposomes; and

sizing the liposomes to a size effective to inhibit formation of precipitated compound, thereby maintaining the entrapped compound in a supersaturated state.